## A. CLASSIFICATION OF SUBJECT MATTER IPC 7 C12Q1/68

C. DOCUMENTS CONSIDERED TO BE RELEVANT

According to International Patent Classification (IPC) or to both national classification and IPC

#### B. FIELDS SEARCHED

Minimum documentation searched (classification system followed by classification symbols) IPC 7 C12Q

Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched

Electronic data base consulted during the international search (name of data base and, where practical, search terms used)

EPO-Internal, WPI Data, PAJ, BIOSIS, EMBASE, CHEM ABS Data, EMBL

Category °	Citation of document, with indication, where appropriate, of the	Relevant to claim No.			
X	PASTINEN T ET AL: "Multiplex, fluorescent, solid-phase minise for efficient screening of DNA variation" CLINICAL CHEMISTRY, AMERICAN AS FOR CLINICAL CHEMISTRY. WINSTON vol. 42, no. 9, 1996, pages 139 XP002126144 ISSN: 0009-9147	sequence SOCIATION , US,	18		
Y	page 1392, left-hand column; ta	ble 1	1-8,19, 20		
X	WO 00/65088 A (AMERSHAM PHARM B ULFENDAHL PER JOHAN (SE); WONG (S) 2 November 2000 (2000-11-02 claims 12,14,21	KIN CHUN	18		
Y	the whole document	-/	1-8,19, 20		
χ Furti	l her documents are listed in the continuation of box C.	Patent family members are listed	n annex.		
'A' docume consid 'E' earlier ( liling d 'L' docume which citatio 'O' docume other ( 'P' docume)	ent defining the general state of the art which is not fared to be of particular relevance document but published on or after the international date ent which may throw doubts on priority claim(s) or is cited to establish the publication date of another n or other special reason (as specified) ent referring to an oral disclosure, use, exhibition or means ent published prior to the international filing date but han the priority date claimed	<ul> <li>'T' later document published after the international filing date or priority date and not in conflict with the application but cled to understand the principle or theory underlying the invention</li> <li>'X' document of particular relevance; the claimed invention cannot be considered novel or cannot be considered to involve an inventive step when the document is taken alone</li> <li>'Y' document of particular relevance; the claimed invention cannot be considered to involve an inventive step when the document is combined with one or more other such documents, such combination being obvious to a person skilled in the art.</li> <li>'&amp;' document member of the same patent family</li> </ul>			
	actual completion of the international search  1 March 2005	Date of mailing of the international sea	•		
Name and r	mailing address of the ISA  European Patent Office, P.B. 5818 Patentlaan 2  NL - 2280 HV Rijswijk  Tel. (+31-70) 340-2040, Tx. 31 651 epo nl, Fax: (+31-70) 340-3016	Authorized officer  Hagenmaier, S			

# PCT/IB2004/004115

ategory °	ation) DOCUMENTS CONSIDERED TO BE RELEVANT  Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No
legory *	Chanch of Cocument, with introducti, where appropriate, of the relevant passages	TOOTER TO ORDING THE
	WORRALL T A ET AL: "Allele-specific	1-8,
	HLA-DR typing by mass spectrometry: an	18-20
	alternative to hybridization-based typing	10 20
	methods."	
	ANALYTICAL CHEMISTRY. 1 NOV 2000,	
	vol. 72, no. 21,	
	1 November 2000 (2000-11-01), pages	
	5233-5238, XP002287583	1
	ISSN: 0003-2700	0 10 10
	the whole document	9,12,13
	LEUSHNER JAMES ET AL: "Automated mass	1-8,
	spectroscopic platform for high throughput	18-20
	DR Beta typing"	
	HUMAN IMMUNOLOGY,	
	vol. 61, no. Supplement 2, 2000, page	
	S126. XP008032510	
	& 26TH ANNUAL MEETING OF THE AMERICAN	
	SOCIETY FOR HISTOCOMPATIBILITY AND	
	IMMUNOGENETICS; LAKE BUENA VISTA, FLORIDA,	
	USA: OCTOBER 10-14, 2000	·
	ISSN: 0198-8859	
	abstract	9,12,13
		1.0
	TOST J ET AL: "GENOTYPING SINGLE	1-8,
	NUCLEOTIDE POLYMORPHISMS BY MASS	18-20
	SPECTROMETRY"	
	MASS SPECTROMETRY REVIEWS, JOHN WILEY AND	
	SONS, NEW YORK, NY, US,	
	vol. 21, no. 6, November 2002 (2002-11),	
	pages 388-418, XP009019382	
	ISSN: 0022-7037	
	the whole document	9,12,13
	TOST JÖRG ET AL: "Molecular haplotyping	1-8,
	at high throughput."	18-20
	NUCLEIC ACIDS RESEARCH. 1 OCT 2002,	10 20
	vol. 30, no. 19, 1 October 2002 (2002-10-01), page e96,	1
	XP002287584	
	ISSN: 1362-4962	0 12 12
	the whole document	9,12,13
	SAUER S ET AL: "EXTENSION OF THE GOOD	1-8,
	ASSAY FOR GENOTYPING SINGLE NUCLEOTIDE	18-20
	POLYMORPHISMS BY MATRIX-ASSISTED LASER	10.20
	DESORPTION/IONIZATION MASS SPECTROMETRY"	
	RAPID COMMUNICATIONS IN MASS SPECTROMETRY,	
	HEYDEN, LONDON, GB,	
	vol. 17, no. 12, 9 May 2003 (2003-05-09),	
	pages 1265-1272, XP009019406	
	ISSN: 0951-4198	
	the whole document	9,12,13
	/	
	I	ı

### PCT/IB2004/004115

Category °	tion) DOCUMENTS CONSIDERED TO BE RELEVANT  Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
ategory *	Cuation of document, with indication, where appropriate, or the tessent bassages	TOOTEN TO OBINTHO
,	SAUER SASCHA ET AL: "Genotyping single-nucleotide polymorphisms by	1-8, 18-20
	<pre>matrix-assisted laser-desorption/ionization time-of-flight mass spectrometry."</pre>	
	JOURNAL OF CHROMATOGRAPHY. B, ANALYTICAL TECHNOLOGIES IN THE BIOMEDICAL AND LIFE SCIENCES. 25 DEC 2002,	
	vol. 782, no. 1-2, 25 December 2002 (2002-12-25), pages 73-87, XP002287585	
4	ISSN: 1570-0232 the whole document	9,12,13
	WO 02/08462 A (LECHNER DORIS; GUT IVO GLYNNE (FR); CT NAT DE GENOTYPAGE (FR))	1-8, 18-20
	31 January 2002 (2002-01-31) the whole document	9,12,13
,	ROZEMULLER: "Reference panels for sequence based typing: Selection criteria for HLA-A and HLA-B" 2000, XP002287586	1-8, 18-20
	ISBN: 0-945278-02-0 Retrieved from the Internet: URL:http://www.ihwg.org/tmanual/TMcontents .htm>	
	'retrieved on 2004-07-05! Chapter 1-B 	9,12,13
	WO 02/18659 A (HAPLOGEN LLC ; LIU XIANGJUN (US)) 7 March 2002 (2002-03-07) the whole document	1-8, 18-20 9,12,13
	US 5 451 512 A (APPLE RAYMOND J ET AL) 19 September 1995 (1995-09-19) the whole document	1-8, 18-20 9,12,13

#### INTERNATIONAL SEARCH REPORT

International application No.

PCT/IB2004/004115

Вох	No. I	Nucleotide and/or amino acid sequence(s) (Continuation of item 1.b of the first sheet)
1.	With	regard to any nucleotide and/or amino acid sequence disclosed in the international application and necessary to the claimed nation, the international search was carried out on the basis of:
•	a.	type of material  X a sequence listing table(s) related to the sequence listing
	b.	format of material  X in written format  X in computer readable form
	c.	time of filing/furnishing  X contained in the international application as filed  X filed together with the international application in computer readable form furnished subsequently to this Authority for the purpose of search
2.		In addition, in the case that more than one version or copy of a sequence listing and/or table relating thereto has been filed or furnished, the required statements that the information in the subsequent or additional copies is identical to that in the application as filed or does not go beyond the application as filed, as appropriate, were furnished.
3.	Addi	tional comments:
· :		

#### INTERNATIONAL SEARCH REPORT

International application No. PCT/IB2004/004115

Box II Observations where certain claims were found unsearchable (Continuation of item 2 of first sheet)
This International Search Report has not been established in respect of certain claims under Article 17(2)(a) for the following reasons:
Claims Nos.: because they relate to subject matter not required to be searched by this Authority, namely:
Claims Nos.:     because they relate to parts of the International Application that do not comply with the prescribed requirements to such an extent that no meaningful International Search can be carried out, specifically:
3. Claims Nos.: because they are dependent claims and are not drafted in accordance with the second and third sentences of Rule 6.4(a).
Box III Observations where unity of invention is lacking (Continuation of Item 3 of first sheet)
This International Searching Authority found multiple inventions in this international application, as follows:
see additional sheet
As all required additional search fees were timely paid by the applicant, this international Search Report covers all searchable claims.
2. As all searchable claims could be searched without effort justifying an additional fee, this Authority did not invite payment of any additional fee.
3. As only some of the required additional search fees were timely paid by the applicant, this International Search Report covers only those claims for which fees were paid, specifically claims Nos.:
4. No required additional search fees were timely paid by the applicant. Consequently, this International Search Report is restricted to the Invention first mentioned in the claims; it is covered by claims Nos.:  claims 1-8, 18-20 (all partially), 9, 12, 13 (completely)
Remark on Protest  The additional search fees were accompanied by the applicant's protest.  No protest accompanied the payment of additional search fees.

#### FURTHER INFORMATION CONTINUED FROM PCT/ISA/ 210

This International Searching Authority found multiple (groups of) inventions in this international application, as follows:

Invention 1 : claims 1-8, 18-20 (all partially), 9,12,13
 (completely)

Method for HLA typing of HLA-A by the unambiguous determination of short DNA sequence elements at positions 98, 414,539,282,571,368,256,292,238 and 270 simultaneously on both parental alleles at a selected number of positions in HLA -A, comprised of the steps for each position a) hybridising a combination of oligonucleotides complementary to all known sequence variants to a DNA strand upstream of a given position b) carrying out a primer extension reaction with at least one of the four dNTP substrates substituted by a terminating analog

c) analysing the products by mass spectrometry, with the resulting masses allowing unambiguous identification of the used primers and added bases; kit for the implementation of such method; use of such method for screening of tissue donors.

Invention 2: 1-8, 18-20 (all partially), 10,14,15 (completely)

Method for HLA typing of HLA-B by the unambiguous determination of short DNA sequence elements at positions 539,419,559,412,272,362,302,363,206 and 369 simultaneously on both parental alleles at a selected number of positions in HLA-B, comprised of the steps for each position

a) hybridising a combination of oligonucleotides complementary to all known sequence variants to a DNA strand upstream of a given position

b) carrying out a primer extension reaction with at least one of the four dNTP substrates substituted by a terminating

c) analysing the products by mass spectrometry, with the resulting masses allowing unambiguous identification of the used primers and added bases; kit for the implementation of such method; use of such method for screening of tissue donors.

Invention 3: claims 1-8, 18-20 (all partially), 11,16,17 (completely)

#### FURTHER INFORMATION CONTINUED FROM PCT/ISA/ 210

Method for HLA typing of HLA-DRB1 by the unambiguous determination of short DNA sequence elements at positions 125,196,197,227,261,286,299,308,341 and 345 simultaneously on both parental alleles at a selected number of positions in HLA-DRB1, comprised of the steps for each position a) hybridising a combination of oligonucleotides complementary to all known sequence variants to a DNA strand upstream of a given position

b) carrying out a primer extension reaction with at least one of the four dNTP substrates substituted by a terminating

analog

c) analysing the products by mass spectrometry, with the resulting masses allowing unambiguous identification of the used primers and added bases; kit for the implementation of such method; use of such method for screening of tissue donors.

Inventions 4-246: claim 21 (partially)

Invention 4:

Use of the primer with Seq.ID 1 to carry out HLA typing. ..ibidem for inventions 5-246, i.e. each of the 242 primers listed in table IV,V and VI.

		Information on patent family members			PCT/IB20041/004115			
	document earch report		Publication date		Patent family member(s)		1	Publication date
WO 00	65088	A	02-11-2000	AU WO	506250 006508			10-11-2000 02-11-2000
WO 020	08462	A	31-01-2002	EP AU CA EP WO US	117621 843510 241720 130363 020846 200405326	1 A 1 A1 8 A1 2 A1	•	30-01-2002 05-02-2002 31-01-2002 23-04-2003 31-01-2002 18-03-2004
WO 02	18659	A	07-03-2002	AU CA CN JP WO US	891770 242107 150198 200452081 021865 200308254	8 A1 2 A 2 T 9 A2		13-03-2002 07-03-2002 02-06-2004 15-07-2004 07-03-2002 01-05-2003
US 54!	51512		19-09-1995	AU BR CA CN EP FI JP NO NZ ZA	274859 920428 208158 107348 054099 92499 806619 92424 24492 920837	0 A 2 A1 4 A 7 A1 9 A 7 A 6 A 4 A		06-05-1993 11-05-1993 06-05-1993 23-06-1993 12-05-1993 06-05-1993 12-03-1996 06-05-1993 26-07-1994 13-05-1993

 $\widetilde{\mathbf{t}}_i$